

EAST Search History

Ref #	Hits	Search Query	DBs	Default Operator	Plurals	Time Stamp
L1	1923	544/235, 514/248	US-PGPUB; USPAT	OR	OFF	2006/08/30 15:25
L2	1361	544/235, 514/248	USPAT	OR	OFF	2006/08/30 15:25
L3	562	544/235, 514/248	US-PGPUB	OR	OFF	2006/08/30 15:25

Day : Wednesday

Date: 8/30/2006
Time: 15:22:06**PALM INTRANET****Inventor Information for 10/799389**

Inventor Name	City	State/Country
ONO, MITSUNORI	LEXINGTON	MASSACHUSETTS
SUN, LIJUN	HARVARD	MASSACHUSETTS
XIA, ZHIQIANG	ACTON	MASSACHUSETTS
KOSTIK, ELENA	ARLINGTON	MASSACHUSETTS
KOYA, KEIZO	CHESTNUT HILL	MASSACHUSETTS
NAGAI, MASAZUMI	LEXINGTON	MASSACHUSETTS
WU, YAMING	LEXINGTON	MASSACHUSETTS

[Appn Info](#)[Contents](#)[Petition Info](#)[Atty/Agent Info](#)[Continuity/Reexam](#)[Foreign](#)

Search Another: Application# or Patent#
 PCT / / or PG PUBS #
 Attorney Docket #
 Bar Code #

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: 10/799,389 Page 3

chain nodes :
10 11 12 16
ring nodes :
1 2 3 4 5 6 7 8 9
ring/chain nodes :
14
chain bonds :
8-16 10-11 11-12 11-14
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9
exact/norm bonds :
1-2 1-6 2-3 3-4 4-5 5-6 6-9 10-11 11-12 11-14
exact bonds :
5-7 7-8 8-9 8-16
isolated ring systems :
containing 1 :

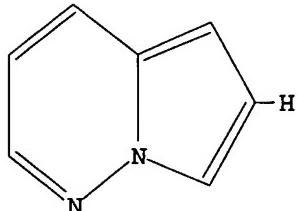
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G2:C,O,S,N,SO2

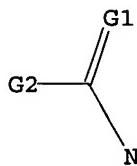
Match level :
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS
11:CLASS 12:CLASS 14:CLASS 16:CLASS

L1 STRUCTURE UPLOADED

=> d 11
L1 HAS NO ANSWERS
L1 STR



G1 O,S,N
G2 C,O,S,N,SO2



Structure attributes must be viewed using STN Express query preparation.

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SAMPLE SCREEN SEARCH COMPLETED - 790 TO ITERATE

100.0% PROCESSED 790 ITERATIONS 6 ANSWERS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**
 PROJECTED ITERATIONS: 14114 TO 17486
 PROJECTED ANSWERS: 6 TO 266

L2 6 SEA SSS SAM L1

=> s 11 sss full
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 FULL SCREEN SEARCH COMPLETED - 16389 TO ITERATE

100.0% PROCESSED 16389 ITERATIONS 75 ANSWERS
 SEARCH TIME: 00.00.01

L3 75 SEA SSS FUL L1

=> file caplus	SINCE FILE	TOTAL
COST IN U.S. DOLLARS	ENTRY	SESSION
FULL ESTIMATED COST	166.94	167.15

FILE 'CAPLUS' ENTERED AT 14:18:35 ON 30 AUG 2006
 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
 PLEASE SEE "HELP USAGETERMS" FOR DETAILS.
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FILE COVERS 1907 - 30 Aug 2006 VOL 145 ISS 10
 FILE LAST UPDATED: 29 Aug 2006 (20060829/ED)

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<http://www.cas.org/infopolicy.html>

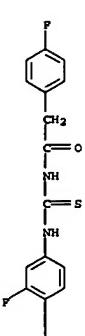
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L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESION NUMBER: 2006-534761 CAPLUS
 DOCUMENT NUMBER: 145:28024
 TITLE: Preparation of fused heterocyclic kinase inhibitors
 INVENTOR(S): Borzilleri, Robert M.; Chen, Zhong; Huynh, Tram N.; Vaccaro, Wayne; Chen, Xiao-Tao; Kim, Kyoung S.; Cai, Zhen-Wei
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 141 pp.
 CODEN: USXKCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

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L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 carboxylate, was given. Compds. I and II inhibit the Met kinase with
 IC50 values between 0.01 to 100 μ M. Pharmaceutical compds. comprising the
 compd. I or II alone or in combination with other antitumor agent are
 disclosed.
 IT 888716-63-0P 888716-64-1P 888716-74-3P
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
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 (preparation of pyrrolopyridines and pyrrolotriazines as kinase
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 for treating cancer)
 RN 888716-63-0 CAPLUS
 CN Benzeneacetamide, 4-fluoro-N-[[3-fluoro-4-(pyrrolo[1,2-b]pyridazin-4-
 yloxy)phenyl]amino]thioxomethyl] - (9CI) (CA INDEX NAME)

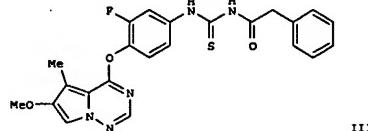
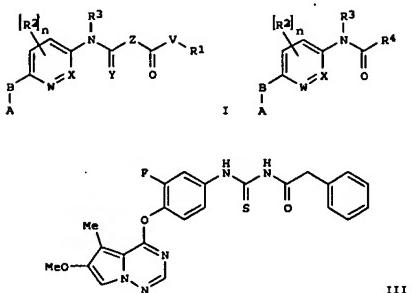


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L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
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 KZ, MD, RU, TJ, TM

PRIORITY APPLN. INFO.: US 2004-583459P P 20040628
 US 2004-612563P P 20040923

OTHER SOURCE(S): MARPAT 145:28024
 GI



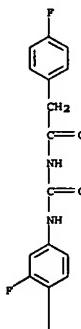
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AB The title compds. I and II [R1 = H, alkyl, cycloalkyl, etc.; R2 = H, halo, CH, etc.; B = O, NR8, S, SO2, CR9C10; V = NR11 or (CR47R48)p; W or X = C or N; Y = O, S, NR12; Z = CR13R14, (CR13R14)mNR15; m = 0-2; n = 0-4; p = 0-4, provided that if p = 0, R1 is not Ph; A = substituted pyrrolo[2,1-f][1,2,4]triazin-4-yl, pyrrolo[1,2-b]pyridazin-4-yl, pyrrolo[2,3-b]pyridin-4-yl, etc.; R3, R8, R11, R15 = H, alkyl, cycloalkyl, etc.; R4 = (un)substituted aryl, heteroaryl, heterocycloalkyl; R9, R10 = H, halo, alkyl, etc.; R12 = H, alkyl, CN, etc.; R13-R15, R47, R48 = H, halo, alkyl, etc.; and their pharmaceutically acceptable salts], useful as protein kinase inhibitors for treating cancer and other protein kinase mediated diseases, were prepared e.g., a multi-step synthesis of III, starting from II.

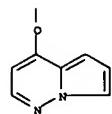
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L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 CN Benzeneacetamide, 4-fluoro-N-[(3-fluoro-4-(pyrrolo[1,2-b]pyridazin-4-
 yloxy)phenyl)amino]carbonyl] - (9CI) (CA INDEX NAME)

PAGE 1-A

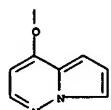


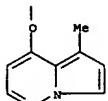
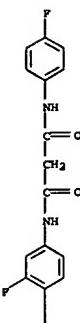
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RN 888716-64-1 CAPLUS
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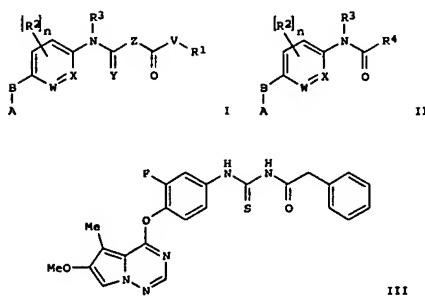




L4 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
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 ZA, ZM, ZW
 RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
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 CG, CI, CM, GA, GN, GO, GM, ML, MR, NE, SN, TD, TG, BW, GH, GM,
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 KZ, MD, RU, TJ, TM

PRIORITY APPLN. INFO.: US 2004-583459P P 20040628
 US 2004-612563P P 20040923

OTHER SOURCE(S): MARPAT 145:28023
 GI



AB: The title compds. I and II [R1 = H, alkyl, cycloalkyl, etc.; R2 = H, halo, etc.; B = O, NR8, S, SO, SO2, CR9C10; V = NR11 or (CR47R48)p; W or X = C or N; Y = O, S, NR12; Z = CR13R14, (CR13R14)mNR15; m = 0-2; n = 0-4; p = 0-4, provided that if p = 0, R1 is not Ph; A = substituted pyrrolo[2,1-f][1,2,4]triazin-4-yl, pyrrolo[1,2-b]pyridazin-4-yl, pyrrolo[2,3-b]pyridin-4-yl, etc.; R3, R8, R11, R12 = H, alkyl, etc.; and their pharmaceutically acceptable salts], useful as protein kinase inhibitors for treating cancer and other protein kinase mediated diseases, were prepared E.g., a multi-step synthesis of III, starting from 5-methyl-4-oxo-3,4-dihydropyrrolo[2,1-f][1,2,4]triazine-6-

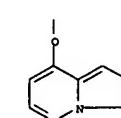
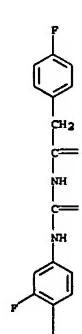
L4 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2006:534671 CAPLUS
 DOCUMENT NUMBER: 145:28023
 TITLE: Preparation of pyrrolopyridines and pyrrolotriazines as kinase inhibitors for treating cancer
 INVENTOR(S): Borzilleri, Robert M.; Chen, Zhong; Hunt, John T.; Huynh, Tram; Poss, Michael A.; Schroeder, Gretchen
 M.; Vaccaro, Wayne; Wong, Tai W.; Chen, Xiao-Tao; Kim, Kyoung S.
 PATENT ASSIGNEE(S): USA
 SOURCE: U.S. Pat. Appl. Publ., 135 pp.
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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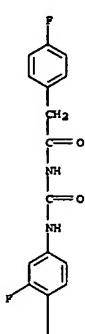
L4 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 carboxylate, was given. Compds. I and II inhibit the Met kinase with IC50 values between 0.01 to 100 μ M. Pharmaceutical compns. comprising the compd. I or II alone or in combination with other antitumor agent are disclosed.

IT 888716-63-0P 888716-64-1P 888716-74-3P
 RW: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (preparation of pyrrolopyridines and pyrrolotriazines as kinase inhibitors for treating cancer)

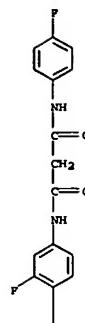
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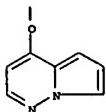
L4 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
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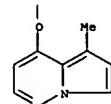
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L4 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 PAGE 1-A

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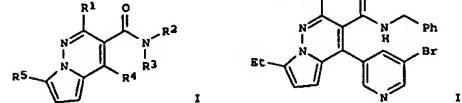
PAGE 2-A



PAGE 2-A

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
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 DOCUMENT NUMBER: 144:128986
 TITLE: Preparation of pyrrolopyridazine derivatives as inhibitors of phosphodiesterase-4 (PDE-IV) and production of tumor necrosis factor- α (TNF- α)
 INVENTOR(S): Abe, Yoshito; Inoue, Makoto; Okumura, Mitsuaki; Ohne, Kazuhiko; Sato, Kentaro
 PATENT ASSIGNEE(S): Astellas Pharma Inc., Japan
 SOURCE: PCT Int. Appl. 52 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006004191	A1	20060112	WO 2005-JP12622	20050701
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RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, PR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BP, BJ, CP, CG, CI, CM, GA, GN, GO, GM, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MM, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
PRIORITY APPLN. INFO.:	AU 2004-903690	A 20040705		
OTHER SOURCE(S):	MARPAT 144:128986			
GI				

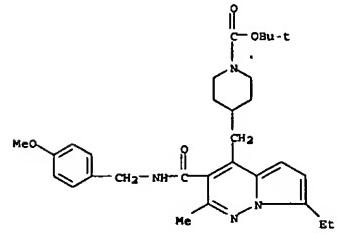


AB Title compds. I [wherein R1 = H, (un)protected carboxy, (un)substituted alkyl, etc.; R2 = (un)substituted (cyclo)alkyl, aryl, etc.; R3 = H or alkyl; R4 and R5 may link together; R4 = (un)substituted (hetero)aryl, etc., and pharmaceutically acceptable salts or prodrugs thereof] were prepared as inhibitors of phosphodiesterase-4 (PDE-IV) and production of tumor necrosis factor- α (TNF- α). For instance, II, which showed inhibition for PDE-IV and on the production of TNF- α with IC50 values of < 1 μ M and 64.0 nM, resp., was synthesized in multiple steps. Therefore, I and their pharmaceutical compns. are useful for the

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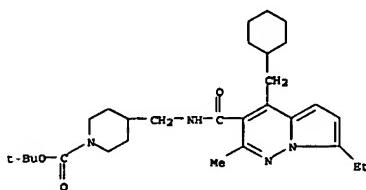
08/30/2006

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 of PDE-IV or TNF- α mediated diseases, such as asthma, COPD and hepatitis.
 IT 873327-04-9P, tert-Butyl 4-[(7-ethyl-3-[(4-methoxybenzyl)amino]carbonyl]-2-methylpyrrolo[1,2-b]pyridazin-4-yl)methyl]-1-piperidinecarboxylate
 RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
 (Inhibitor; preparation of pyrrolopyridazine derivs. as inhibitors of phosphodiesterase-4 and production of tumor necrosis factor- α)
 RN 873327-04-9 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[(7-ethyl-3-[(4-methoxybenzyl)amino]carbonyl]-2-methylpyrrolo[1,2-b]pyridazin-4-yl)methyl]-, 1,1-dimethylallyl ester (9CI) (CA INDEX NAME)

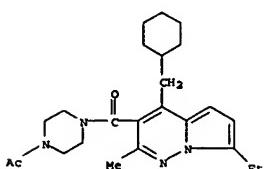


IT 873327-01-6P, tert-Butyl 4-[[[(4-(cyclohexylmethyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)carbonyl]amino]methyl]-1-piperidinecarboxylate 873327-03-8P, 3-[(4-Acetyl-1-piperazinyl)carbonyl]-4-(cyclohexylmethyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl-10-7P, 3-[(4-Acetyl-1-piperazinyl)carbonyl]-4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl-12-9B, 7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl-13-10P, tert-Butyl 4-[[[(4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl)carbonyl]amino]methyl]-1-piperidinecarboxylate 873327-12-9B, tert-Butyl 4-[[[(4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl)carbonyl]amino]methyl]-1-piperidinecarboxylate 873327-13-0P, tert-Butyl 4-[[[(4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl)carbonyl]amino]methyl]-1-piperidinecarboxylate 873327-37-8P, 4-[(1-Acetyl-4-piperidinyl)methyl]-7-ethyl-N-(4-methoxybenzyl)-2-methylpyrrolo[1,2-b]pyridazin-3-carboxamide 873327-48-1P, tert-Butyl 4-[[[(4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl)carbonyl]amino]methyl]-1-piperidinecarboxylate 873327-49-2P, tert-Butyl 4-[[[(7-ethyl-2-methyl-4-(tetrahydro-2H-pyran-4-yl)methyl)pyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino]methyl]-1-piperidinecarboxylate 873327-50-5P, tert-Butyl 4-[(7-ethyl-2-methyl-4-(tetrahydro-2H-pyran-4-yl)methyl)pyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino]methyl]-1-piperidinecarboxylate
 RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (Inhibitor; preparation of pyrrolopyridazine derivs. as inhibitors of

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 phosphodiesterase-4 and prodn. of tumor necrosis factor- α)
 RN 873327-01-6 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[[[4-(cyclohexylmethyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino)methyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



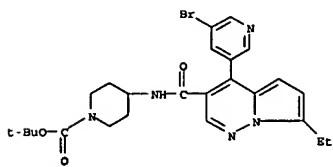
RN 873327-03-8 CAPLUS
 CN Piperazine, 1-acetyl-4-[(4-(cyclohexylmethyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)carbonyl]- (9CI) (CA INDEX NAME)



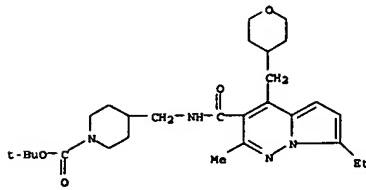
RN 873327-10-7 CAPLUS
 CN Piperazine, 1-acetyl-4-[(4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl)carbonyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)

RN 873327-48-1 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[[[4-(5-bromo-3-pyridinyl)-7-ethylpyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



RN 873327-49-2 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[[[7-ethyl-2-methyl-4-[(tetrahydro-2H-pyran-4-yl)methyl]pyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino)methyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)



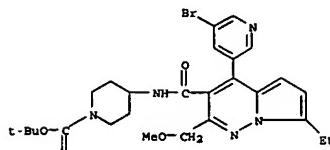
RN 873327-50-5 CAPLUS

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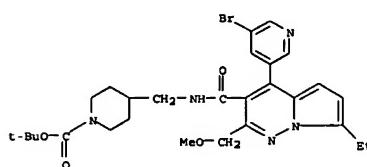
L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



RN 873327-12-9 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[[[4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

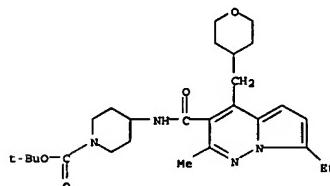


RN 873327-13-0 CAPLUS
 CN 1-Piperidinecarboxylic acid, 4-[[[4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino)methyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

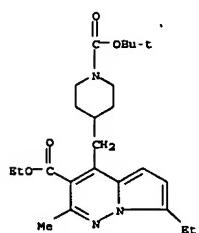


RN 873327-37-8 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-carboxamide, 4-[(1-acetyl-4-piperidinyl)methyl]-7-ethyl-N-[(4-methoxyphenyl)methyl]-2-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
 1-Piperidinecarboxylic acid, 4-[[[7-ethyl-2-methyl-4-[(tetrahydro-2H-pyran-4-yl)methyl]pyrrolo[1,2-b]pyridazin-3-yl]carbonyl]amino]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

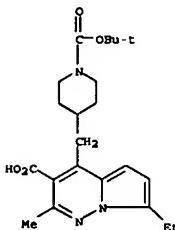


IT 873326-82-0P, Ethyl 4-[[1-(tert-butoxycarbonyl)-4-piperidinyl]methyl]-7-ethyl-2-methylpyrrolo[1,2-b]pyridazine-3-carboxylate
 873326-89-7P, 4-[[1-(tert-Butoxycarbonyl)-4-piperidinyl]methyl]-7-ethyl-2-methylpyrrolo[1,2-b]pyridazine-3-carboxylic acid
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation of pyrrolopyridazine derivs. as inhibitors of phosphodiesterase-4 and production of tumor necrosis factor- α)
 RN 873326-82-0 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-carboxylic acid, 4-[[1-((1,1-dimethylethoxy)carbonyl)-4-piperidinyl)methyl]-7-ethyl-2-methyl-, ethyl ester (9CI) (CA INDEX NAME)



RN 873326-89-7 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-carboxylic acid, 4-[[1-((1,1-dimethylethoxy)carbonyl)-4-piperidinyl)methyl]-7-ethyl-2-methyl- (9CI) (CA INDEX NAME)

L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2004:799444 CAPLUS
DOCUMENT NUMBER: 141:296005
TITLE: Preparation of fused pyrrole compounds as TNF α and/or PDE4 inhibitors for treatment of cancer, inflammatory disorders, and autoimmune diseases

INVENTOR(S): Omo, Mitsuori; Sun, Lijun; Xia, Zhi Qiang; Kostik, Elena; Koya, Keizo; Wu, Yaming; Nagai, Masazumi

PATENT ASSIGNEE(S): Synta Pharmaceuticals Corp., USA

SOURCE: PCT Int. Appl., 86 pp.

CODEN: PIIXD3

DOCUMENT TYPE: Patent

LANGUAGE: English

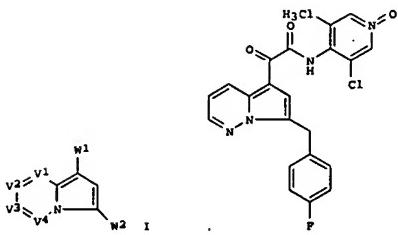
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004082606	A2	20040930	WO 2004-US7469	20040311
WO 2004082606	A3	20050127		
WO 2004082606	C1	20050303		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PR, PL, PT, RU, SC, SD, SE, SO, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RN: BW, GH, GM, KB, LS, MM, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, PI, PR, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CO, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2004222387	A1	20040930	AU 2004-222387	20040311
CA 2517034	AA	20040930	CA 2004-2517034	20040311
EP 1601678	A2	20051207	EP 2004-719768	20040311
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IS, SI, LT, LV, PI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK				
US 2005014754	A1	20050120	US 2004-799389	20040312
PRIORITY APPLN. INFO.: US 2005014754 P 20030312				
WO 2004-US7469 A 20040311				

OTHER SOURCE(S): MARPAT 141:296005
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L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



AB Title compds. I [wherein one of W1 and W2 = YC(-Z)NR1R2 and the other = XR3; V1-V4 = independently N, (un)substituted CH; or V1V2 or V3V4 = S, O, (un)substituted NH; or 2 adjacent V's form a fused aryl ring; X = bond, O, S, SO, SO2, CO, (un)substituted CH2, NH, CONH, NHCO; Y = O, S, SO, SO2, CO, (un)substituted CH2, NH, CONH, NHCO; Z = O, S, (un)substituted NOH, NH; R1, R2 = independently H, (un)substituted aliphatic group, heterocyclyl, aryl; or NR1R2 = heterocyclyl, heteroaryl; R3 = (un)substituted aryl, aliphatic; with provisos; and pharmaceutically acceptable salts and prodrugs thereof], such as indolizines, pyrrolo[1,2-b]pyridazines, and pyrrolo[2,1-b]thiazoles, were prepared as phosphodiesterase IV (PDE4) and/or tumor necrosis factor α (TNF α) inhibitors. For example, reaction of 2-bromo-4'-fluoroacetophenone with 3-methylpyridazine in the presence of DMF-Me2SO4 provided (4-fluorophenyl)(pyrrolo[1,2-b]pyridazin-7-yl)methanone, which was reduced to the benzyl derivative using BH3-THF. Sequential coupling with oxalyl chloride and N-oxide gave II (1.5% overall). The latter inhibited TNF α in human peripheral blood cells and PDE4 in U937 human monocytic cells with IC50 values of about 50 nM and about 5 nM, resp. II also demonstrated in vitro anticancer activity in human cancer cell line MDA435 with an IC50 value of about 1 μ M. Thus, I and their pharmaceutical compns. are useful for the treatment or prevention of cancer, inflammatory disorders, autoimmune diseases, and other conditions involving PDE4 or elevated levels of cytokines.

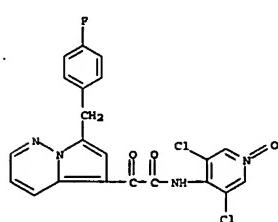
IT 764723-23-1P, N-(3,5-Dichloro-1-oxypyridin-4-yl)-2-[7-(4-fluorobenzyl)pyrrolo[1,2-b]pyridazin-5-yl]-2-(oxo)acetamide 764723-27-5P, 2-[7-(4-Cyanobenzyl)pyrrolo[1,2-b]pyridazin-5-yl]-N-(3,5-dichloropyridin-4-yl)-2-(oxo)acetamide 764723-28-6P, 2-[7-(4-Methoxybenzyl)pyrrolo[1,2-b]pyridazin-5-yl]-2-(oxo)-N-(pyridin-4-yl)acetamide 764723-29-7P, 2-[7-(4-Chlorobenzyl)pyrrolo[1,2-b]pyridazin-5-yl]-N-(isoxazol-5-yl)-2-(oxo)acetamide

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU

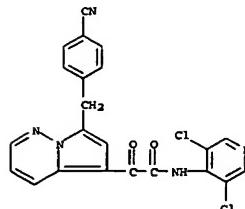
L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(cytokine inhibitor; prepn. of fused pyrrole compds. as TNF α and/or PDE4 inhibitors for treatment of cancer, inflammatory disorders, and autoimmune diseases)

RN 764723-23-1 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-5-acetamide, N-(3,5-dichloro-1-oxido-4-pyridinyl)-7-[(4-fluorophenyl)methyl]-^aoxo- (9CI) (CA INDEX NAME)



RN 764723-27-5 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-5-acetamide, 7-[(4-cyanophenyl)methyl]-N-(3,5-dichloro-4-pyridinyl)-^aoxo- (9CI) (CA INDEX NAME)

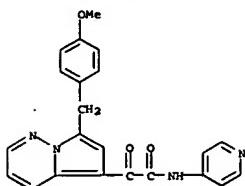


RN 764723-28-6 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-5-acetamide, 7-[(4-methoxyphenyl)methyl]-^aoxo-N-4-pyridinyl- (9CI) (CA INDEX NAME)

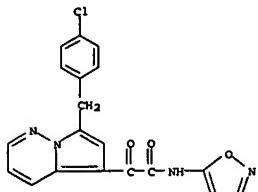
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work

L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

(Continued)



RN 764723-29-7 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-5-acetamide, 7-[(4-chlorophenyl)methyl]-N-5-isoxazolyl-a-oxo- (9CI) (CA INDEX NAME)



L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:606471 CAPLUS

DOCUMENT NUMBER: 141:157123

TITLE: Preparation of pyrrolopyridazines as inhibitors of phosphodiesterase IV (PDE IV) and production of tumor necrosis factor- α (TNF- α)

INVENTOR(S): Abe, Yoshito; Inoue, Makoto; Mizutani, Teiyoshi; Sawada, Kozo; Ohne, Kazuhiko; Okumura, Mitsuaki; Sawada, Yukio; Imamura, Kenichiro

PATENT ASSIGNEE(S): Fujisawa Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 360 pp.

CODEN: PIXKD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

WO 2004063197	A1	20040729	WO 2003-JP17091	20031226
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HU, ID, IL, IN, IS, JP, KB, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MM, MX, MZ, NI, NO, NZ, OM, PO, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, KE, LS, MM, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BV, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CO, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD,				

TG CA 2513081 AA 20040729 CA 2003-2513081 20031226

AU 2003294183 A1 20040810 AU 2003-294183 20031226

EP 15815535 A1 20051005 EP 2003-789642 20031226

R: AT, BE, CH, DE, DK, ES, PR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK

BR 2003017358 A 20051213 BR 2003-17358 20031226

CN 1759117 A 20060412 CN 2003-80110135 20031226

JP 2006515597 T2 20060601 JP 2004-566309 20031226

US 2005075342 A1 20050407 US 2003-747079 20031226

NO 2005003748 A 20051007 NO 2005-3748 20050804

PRIORITY APPLN. INFO.: AU 2003-900189 A 20031019

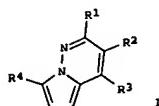
AU 2003-903628 A 20030714

WO 2003-JP17091 W 20031226

OTHER SOURCE(S): MARPAT 141:157123

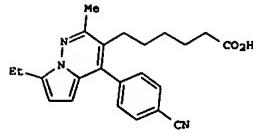
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L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



RN 728011-71-0 CAPLUS

CN Pyrrolo[1,2-b]pyridazine-3-carboxylic acid, 7-ethyl-2-methyl-4-[4-((phenylmethoxy)carbonyl)amino]sulfonylphenyl-, ethyl ester (9CI) (CA INDEX NAME)



AB Title compds. I [wherein R₁ = (un)protected CO₂H, CONH₂ and derive., OH and lower alkoxy, mono/di/cyclo(lower)alkylamino, trihalo(lower)alkyl, (un)substituted lower alkyl, aryl, heterocyclyl; R₂ = R₇, or

- (A)1-p-XA2A7;

p = 0-1; A1 = ethylene, HC:CH; A2 = (CH₂)_n, (HC:CH)_m; n = 1-6; m = 1-3; X = a single bond, O, NH and derive., C(=O), hydroxalkyne, etc.; R₇ = H, (un)substituted aryl, heterocyclyl, CO₂H and derive., acyl, CN, NH₂ and derive., OH, aryloxy, acyloxy; R1R2 = (un)substituted lower alk(en)ylene, optionally interrupted by NH₂ or sulfonyl, and optionally fused with benzene; R₃ = (un)substituted aryl, heterocyclyl; R₄ = H, halo, CN, carbamoyl, acyl, thiocyanate, lower alkylthio, lower alk(en)yl, hydroxy(lower)alkyl, trihalo(lower)alkyl; and their pharmaceutically acceptable salts or prodrugs were prepared as inhibitors of phosphodiesterase IV (PDE IV) and production of tumor necrosis factor- α (TNF- α). Thus, reacting Et 7-(4-cyanobenzoyl)-8-oxononanoate (preparation given) with 2-ethyl-1H-pyrrol-1-amine in toluene in the presence

of p-TSA at reflux, followed ester hydrolysis in the presence of KOH/MeOH gave pyrrolopyridazine II and its 4-(aminocarbonyl)phenyl derivative. Pyrrolopyridazine II displayed an IC₅₀ < 1 μ M for PDE IV inhibition.

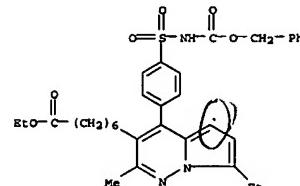
II gave an IC₅₀ < 100 μ M for the inhibition of TNF- α production I are useful for treating asthma, COPD, fibrosis, hepatitis, Alzheimer's diseases, etc.

IT 728011-71-0P, Ethyl 7-[4-(4-[(benzyl oxy)carbonyl]amino)sulfonyl]phenyl-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-ylheptanoate 728015-23-4P, Ethyl 2-(2-amino-2-oxoethyl)-4-(4-cyanophenyl)-7-ethylpyrrolo[1,2-b]pyridazine-3-carboxylate 728015-80-3P, text-Butyl

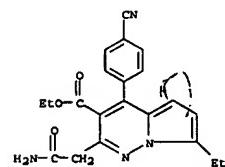
[4-(4-(3-cyanophenyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)butyl]carbamate 728017-66-1P, Ethyl 5-[4-(5-(text-

butoxycarbonyl)amino]-3-pyridinyl]-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl]pentanoate

Habte



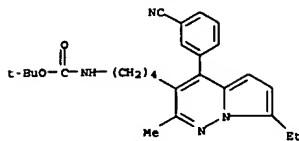
RN 728015-23-4 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-carboxylic acid, 2-(2-amino-2-oxoethyl)-4-(4-cyanophenyl)-7-ethyl-, ethyl ester (9CI) (CA INDEX NAME)



RN 728015-80-3 CAPLUS
 CN Carbamic acid, [4-(4-(3-cyanophenyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)butyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

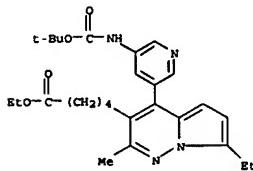
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

(Continued)



RN 728017-66-1 CAPLUS

CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-[5-[(1,1-dimethylethoxy)carbonyl]amino]-3-pyridinyl]-7-ethyl-2-methyl-, ethyl ester (9CI) (CA INDEX NAME)



IT 728015-56-3P, 2-[3-(4-(3-Chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)pentanoyl]methanesulfonamide

728015-59-6P 728015-68-7P 728015-72-3P, N-[5-(4-(3-Cyanophenyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)pentanoyl]methanesulfonamide 728016-38-4P, 5-(4-(3-Cyanophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)pentanamide 728016-40-8P, 5-(4-(3-Chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)N-(2-pyridinyl)pentanamide 728016-44-2P, N-[5-(4-(3-Chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)pentanoyl]methanesulfonamide 728016-44-2P,

5-[4-(2-Chloro-4-pyridinyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl]-N-(2-pyridinyl)pentanamide 728016-46-4P, N-[5-(4-(5-Bromo-3-pyridinyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)pentanoyl]methanesulfonamide 728016-66-8P, 5-(4-(3-Cyanophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)-N-(2-pyridinyl)pentanamide 728016-68-0P, N-[5-(4-(3-Chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)pentanoyl]methanesulfonamide 728016-70-4P, 5-[4-(3-Chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl]N-methylpentanamide 728016-72-6P,

5-[4-(2-Chloro-4-pyridinyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl]-

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)
Ethyl 5-[2-[(2-(benzylamino)-2-oxethoxy)methyl]-7-ethyl-4-(5-methyl-3-pyridinyl)pyrrolo[1,2-b]pyridazin-3-yl]carbamate 728021-59-8P, tert-Butyl [2-[(5-bromo-3-pyridinyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl]pentanoyl]carbamate 728021-66-7P

728021-67-8P, Ethyl 5-[4-(5-bromo-3-pyridinyl)-2-[(dimethylamino)carbonyloxy]methyl]-7-ethylpyrrolo[1,2-b]pyridazin-3-yl]pentanone 728021-68-9P, 5-[4-(5-Bromo-3-pyridinyl)-3-(5-ethoxy-5-oxopentyl)-2-phenylpyrrolo[1,2-b]pyridazine-2-yl]methyl 1-pyrrolidinecarboxylate 728021-69-0P, Ethyl

5-[4-(5-bromo-3-pyridinyl)-7-ethyl-2-[[[(methylphenyl)amino]carbonyloxy]methyl]-2-phenylpyrrolo[1,2-b]pyridazin-3-yl]pentanone 728021-70-3P, [4-(5-Bromo-3-pyridinyl)-3-(4-ethoxy-4-oxobutyl)-7-ethylpyrrolo[1,2-b]pyridazin-2-yl]methyl 4-morpholinocarboxylate 728021-72-5P, Ethyl

4-[4-(5-bromo-3-pyridinyl)-2-[[[(dimethylamino)carbonyloxy]methyl]-7-ethylpyrrolo[1,2-b]pyridazin-3-yl]butanate 728021-74-7P, Ethyl

3-[4-(5-bromo-3-pyridinyl)-2-[[[(dimethylamino)carbonyloxy]methyl]-7-ethylpyrrolo[1,2-b]pyridazin-3-yl]propanoate

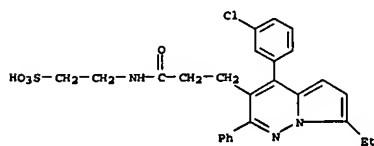
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(phosphodiesterase IV inhibitor; prepn. of pyrrolopyridazines as inhibitors of phosphodiesterase IV and prodn. of tumor necrosis factor- α (TNF- α))

RN 728015-56-3 CAPLUS

CN Ethanesulfonic acid,

2-[3-(4-(3-chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)-1-oxopropyl]amino] (9CI) (CA INDEX NAME)



RN 728015-59-6 CAPLUS

CN Pyrrolo[1,2-b]pyridazine-3-propanamide, 4-(3-chlorophenyl)-7-ethyl-2-phenyl-N-[2,3,4,6-tetrakis-O-(2,2-dimethyl-1-oxopropyl)-D-galactopyranosyl]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

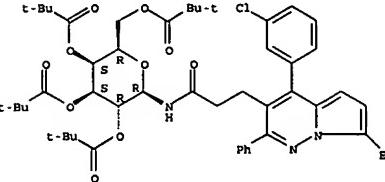
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)

N-methylpentanamide 728016-74-8P, 3-[7-Ethyl-3-[5-(4-morpholinyl)-5-oxopentyl]-2-phenylpyrrolo[1,2-b]pyridazin-4-yl]benzonitrile 728016-76-0P, 3-[7-Ethyl-2-methyl-3-[5-(4-morpholinyl)-5-oxopentyl]pyrrolo[1,2-b]pyridazin-4-yl]benzonitrile 728016-78-2P, 5-[4-(3-Cyanophenyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl]pentanamide 728017-85-4P, Ethyl 5-[7-ethyl-4-(5-methyl-3-pyridinyl)-2-[(2-(4-morpholinyl)-2-oxethoxy)methyl]pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728017-86-5P, Ethyl 5-[7-ethyl-2-[(2-(methylamino)-2-oxethoxy)methyl]pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728017-90-1P, Ethyl 5-[2-[(acetylamino)methyl]-7-ethyl-4-(5-methyl-3-pyridinyl)pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728017-94-5P, Ethyl 5-[7-ethyl-2-[(methoxycarbonyl)amino)methyl]-4-(5-methyl-3-pyridinyl)pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728018-21-1P, 5-[7-Ethyl-4-(5-methyl-3-pyridinyl)-2-[(2-(4-morpholinyl)-2-oxethoxy)methyl]pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728018-22-2P, 5-[7-Ethyl-2-[(2-(methylamino)-2-oxethoxy)methyl]pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728018-30-2P, 5-[2-[(Acetylamino)methyl]-7-ethyl-4-(5-methyl-3-pyridinyl)pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728018-34-6P, Ethyl 5-[7-Ethyl-2-[(methoxycarbonyl)amino)methyl]-4-(5-methyl-3-pyridinyl)pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728018-46-0P, 5-[4-(Bromo-3-pyridinyl)-7-ethyl-2-methyl-3-[3-(4-morpholinyl)-3-oxopropyl]pyrrolo[1,2-b]pyridazine 728018-47-1P,

3-[4-(5-Bromo-3-pyridinyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl]-N-methylpropanamide 728018-48-2P, N-[3-(4-(5-Bromo-3-pyridinyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)propanoyl]methanesulfonamide 728018-49-3P, 2-[3-(4-(3-Chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)propanoyl]methanesulfonic acid 728020-31-3P, 5-[4-(5-Bromo-3-pyridinyl)-7-ethyl-2-[(4-morpholinyl)carbonyloxy]methyl]pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728020-32-4P, 5-[4-(5-Bromo-3-pyridinyl)-2-[(dimethylamino)carbonyloxy]methyl]pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728020-35-7P,

4-[4-(5-Bromo-3-pyridinyl)-7-ethyl-2-[(4-morpholinyl)carbonyloxy]methyl]pyrrolo[1,2-b]pyridazin-3-yl]butanoic acid 728020-36-8P, 4-[4-(5-Bromo-3-pyridinyl)-2-[(dimethylamino)carbonyloxy]methyl]-7-ethylpyrrolo[1,2-b]pyridazin-3-yl]butanoic acid 728020-37-9P, 3-[4-(5-Bromo-3-pyridinyl)-2-[(dimethylamino)carbonyloxy]methyl]-7-ethylpyrrolo[1,2-b]pyridazin-3-yl]butanoic acid 728020-57-3P, 5-[2-[(2-Benzylamino)-2-oxethoxy)methyl]-7-ethyl-4-(5-methyl-3-pyridinyl)pyrrolo[1,2-b]pyridazin-3-yl]pentanone 728020-98-2P, 728021-11-2P, 728021-13-4P, N-[2-Aminoethyl]-3-[4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl]propanamide 728021-52-1P, 3-[4-(5-Bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl]N-[2-hydroxy-1,1-bis(hydroxymethyl)ethyl]propanamide 728021-53-2P, tert-Butyl [2-[(4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)pyrrolo[1,2-b]pyridazin-3-yl)propanoyl]amino]ethyl carbamate 728021-54-3P,

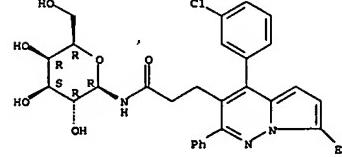
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



RN 728015-68-7 CAPLUS

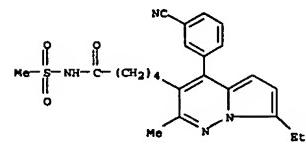
CN Pyrrolo[1,2-b]pyridazine-3-propanamide, 4-(3-chlorophenyl)-7-ethyl-N-P-D-galactopyranosyl-2-phenyl- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 728015-72-3 CAPLUS

CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-cyanophenyl)-7-ethyl-2-methyl-N-(methylsulfonyl)- (9CI) (CA INDEX NAME)

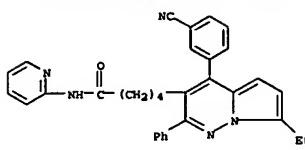


RN 728016-38-4 CAPLUS

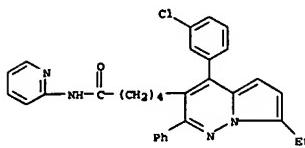
CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-cyanophenyl)-7-ethyl-2-phenyl-N-2-pyridinyl- (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

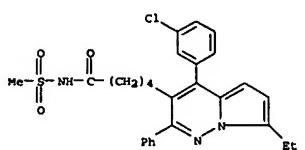
(Continued)



RN 728016-40-8 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-chlorophenyl)-7-ethyl-2-phenyl-N-2-pyridinyl- (9CI) (CA INDEX NAME)



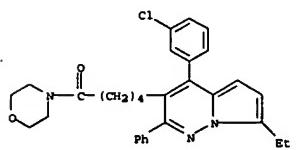
RN 728016-42-0 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-chlorophenyl)-7-ethyl-N-(methylsulfonyl)-2-phenyl- (9CI) (CA INDEX NAME)



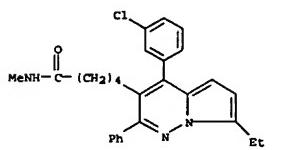
RN 728016-44-2 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(2-chloro-4-pyridinyl)-7-ethyl-2-phenyl-N-2-pyridinyl- (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

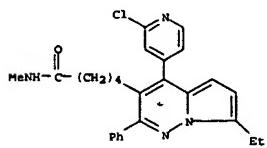
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RN 728016-70-4 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-chlorophenyl)-7-ethyl-N-methyl-2-phenyl- (9CI) (CA INDEX NAME)



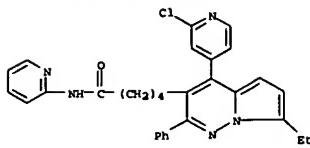
RN 728016-72-6 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(2-chloro-4-pyridinyl)-7-ethyl-N-methyl-2-phenyl- (9CI) (CA INDEX NAME)



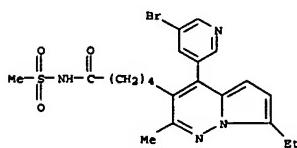
RN 728016-74-8 CAPLUS
 CN Morpholine, 4-[5-(4-(3-cyanophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)-1-oxopentyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

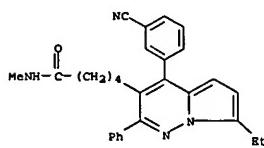
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RN 728016-46-4 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-methyl-N-(methylsulfonyl)- (9CI) (CA INDEX NAME)



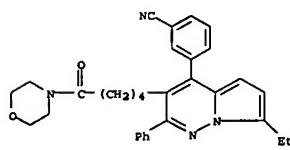
RN 728016-66-8 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-cyanophenyl)-7-ethyl-N-methyl-2-phenyl- (9CI) (CA INDEX NAME)



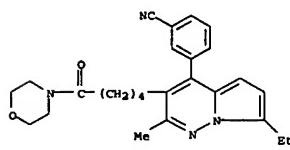
RN 728016-68-0 CAPLUS
 CN Morpholine, 4-[5-(4-(3-chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl)-1-oxopentyl]- (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

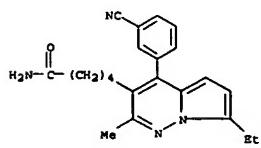
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RN 728016-76-0 CAPLUS
 CN Morpholine, 4-[5-(4-(3-cyanophenyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)-1-oxopentyl]- (9CI) (CA INDEX NAME)



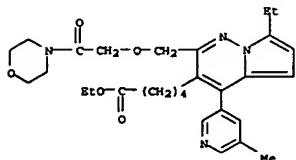
RN 728016-78-2 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanamide, 4-(3-cyanophenyl)-7-ethyl-2-methyl- (9CI) (CA INDEX NAME)



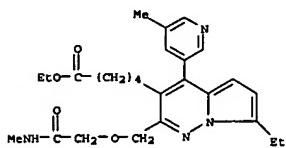
RN 728017-85-4 CAPLUS
 CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-4-(5-methyl-3-pyridinyl)-2-[(2-(4-morpholinyl)-2-oxoethoxy)methyl]-, ethyl ester (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

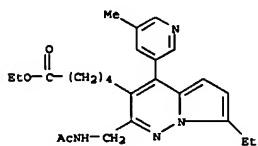
(Continued)



RN 728017-86-5 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-2-[(2-(methylamino)-2-oxoethoxy)methyl]-4-(5-methyl-3-pyridinyl)-, ethyl ester (9CI) (CA INDEX NAME)



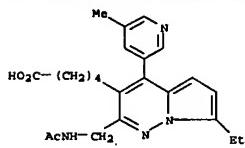
RN 728017-90-1 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 2-[(acetylamino)methyl]-7-ethyl-4-(5-methyl-3-pyridinyl)-, ethyl ester (9CI) (CA INDEX NAME)



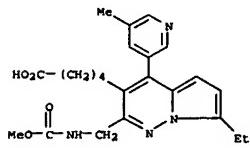
RN 728017-94-5 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-2-[(methoxycarbonyl)amino)methyl]-4-(5-methyl-3-pyridinyl)-, ethyl ester (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

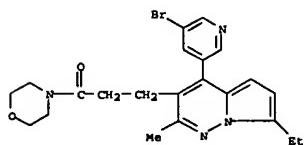
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RN 728018-34-6 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-2-[(methoxycarbonyl)amino)methyl]-4-(5-methyl-3-pyridinyl)- (9CI) (CA INDEX NAME)



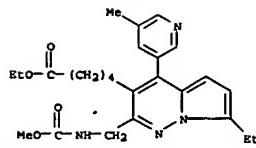
RN 728018-46-0 CAPLUS
CN Morpholine, 4-[(3-[(5-bromo-3-pyridinyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl]-1-oxopropyl)- (9CI) (CA INDEX NAME)



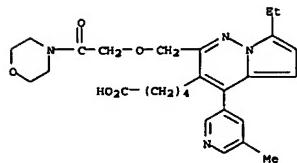
RN 728018-47-1 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanamide, 4-(5-bromo-3-pyridinyl)-7-ethyl-N,2-dimethyl- (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

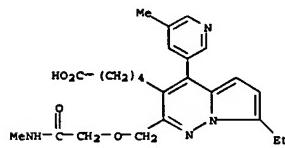
(Continued)



RN 728018-21-1 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-4-(5-methyl-3-pyridinyl)-2-[(2-(4-morpholinyl)-2-oxoethoxy)methyl]- (9CI) (CA INDEX NAME)



RN 728018-22-2 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-2-[(2-(methylamino)-2-oxoethoxy)methyl]-4-(5-methyl-3-pyridinyl)- (9CI) (CA INDEX NAME)



RN 728018-30-2 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 2-[(acetylamino)methyl]-7-ethyl-4-(5-methyl-3-pyridinyl)- (9CI) (CA INDEX NAME)

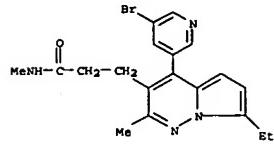
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

(Continued)

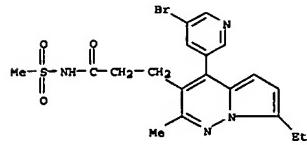


L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN

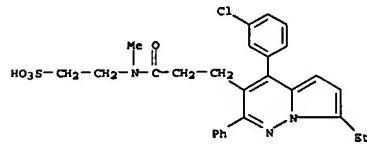
(Continued)



RN 728018-48-2 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanamide, 4-[(3-[(5-bromo-3-pyridinyl)-7-ethyl-2-methyl-N-(methylsulfonyl)- (9CI) (CA INDEX NAME)



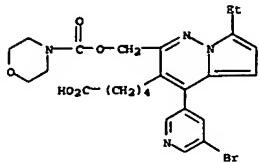
RN 728018-49-3 CAPLUS
CN Ethanesulfonic acid, 2-[(3-[(4-(3-chlorophenyl)-7-ethyl-2-phenylpyrrolo[1,2-b]pyridazin-3-yl]-1-oxopropyl)methylamino]- (9CI) (CA INDEX NAME)



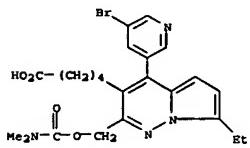
RN 728020-31-3 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(4-morpholinylcarbonyl)oxy)methyl]- (9CI) (CA INDEX NAME)

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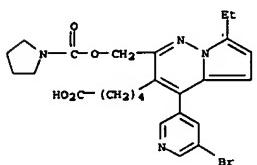
(Continued)



RN 728020-32-4 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-2-[(dimethylamino)carbonyloxy]methyl- (9CI) (CA INDEX NAME)



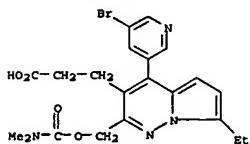
RN 728020-33-5 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(1-pyrrolidinylcarbonyl)oxy]methyl- (9CI) (CA INDEX NAME)



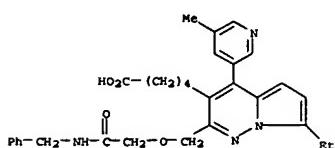
RN 728020-34-6 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(methylphenylamino)carbonyl]oxy]methyl- (9CI) (CA INDEX NAME)

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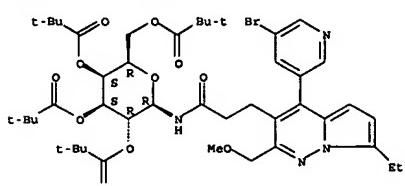
(Continued)



RN 728020-57-3 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-4-(5-methyl-3-pyridinyl)-2-[(2-oxo-2-[(phenylmethoxy)amino]ethoxy)methyl]- (9CI) (CA INDEX NAME)



Absolute stereochemistry.



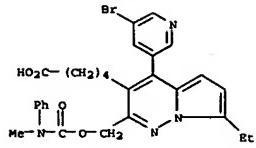
RN 728021-11-2 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanamide, 4-(5-bromo-3-pyridinyl)-7-ethyl-N-[2-(3,4,6-tetrahydro-2H-1,3-dimethyl-1-oxopropyl)-β-D-galactopyranosyl]-3-(methoxymethyl)- (9CI) (CA INDEX NAME)

Habte

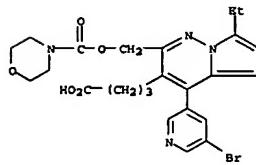
08/30/2006

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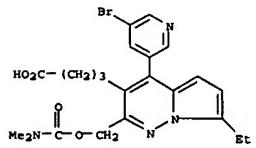
(Continued)



RN 728020-35-7 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-butanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(4-morpholinylcarbonyl)oxy]methyl- (9CI) (CA INDEX NAME)



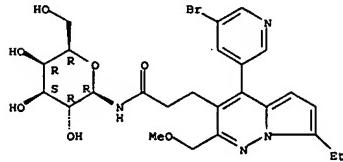
RN 728020-36-8 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-butanoic acid, 4-(5-bromo-3-pyridinyl)-2-[(dimethylamino)carbonyl]oxy]methyl- (9CI) (CA INDEX NAME)



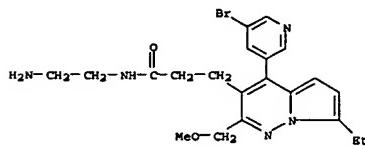
RN 728020-37-9 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanoic acid, 4-(5-bromo-3-pyridinyl)-2-[(2-hydroxy-2-[(phenylmethoxy)amino]ethoxy)methyl]- (9CI) (CA INDEX NAME)

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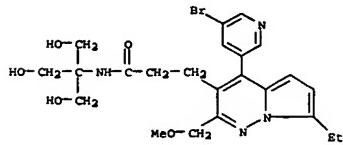
(Continued)



RN 728021-13-4 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanamide, N-(2-aminoethyl)-4-(5-bromo-3-pyridinyl)-7-ethyl-2-(methoxymethyl)- (9CI) (CA INDEX NAME)

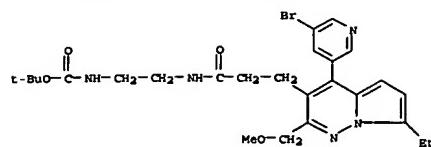


RN 728021-52-1 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanamide, 4-(5-bromo-3-pyridinyl)-7-ethyl-N-[2-hydroxy-1,1-bis(hydroxymethyl)ethyl]- (9CI) (CA INDEX NAME)

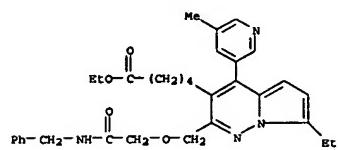


RN 728021-53-2 CAPLUS
CN Carbamic acid, 2-[(3-[(4-(5-bromo-3-pyridinyl)-7-ethyl-1-(methoxymethyl)amino)ethyl]-1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



RN 728021-54-3 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 7-ethyl-4-(5-methyl-3-pyridinyl)-2-[(2-oxo-2-[(phenylmethyl)amino]ethoxy)methyl]-, ethyl ester (9CI) (CA INDEX NAME)

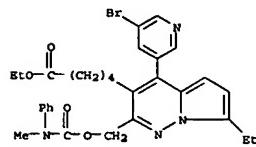


RN 728021-59-8 CAPLUS
CN Carbamic acid, [2-(4-(5-bromo-3-pyridinyl)-7-ethyl-2-methylpyrrolo[1,2-b]pyridazin-3-yl)ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

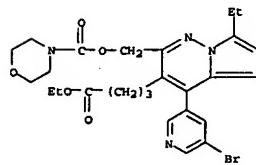


RN 728021-66-7 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(4-morpholinylcarbonyl)oxy]methyl-, ethyl ester (9CI) (CA INDEX NAME)

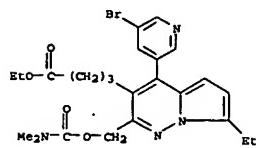
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



RN 728021-70-3 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-butanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(4-morpholinylcarbonyl)oxy]methyl-, ethyl ester (9CI) (CA INDEX NAME)

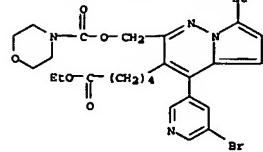


RN 728021-72-5 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-butanoic acid, 4-(5-bromo-3-pyridinyl)-2-[(dimethylamino)carbonyl]oxy]methyl-, ethyl ester (9CI) (CA INDEX NAME)

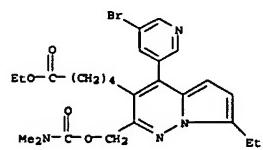


RN 728021-74-7 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-propanoic acid, 4-(5-bromo-3-pyridinyl)-2-[(dimethylamino)carbonyl]oxy]methyl-, ethyl ester (9CI) (CA INDEX NAME)

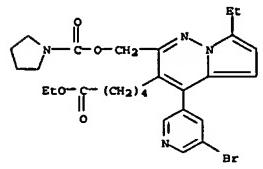
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



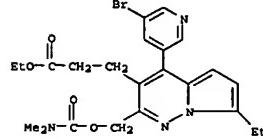
RN 728021-67-8 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-2-[(dimethylamino)carbonyl]oxy]methyl-, ethyl ester (9CI) (CA INDEX NAME)



RN 728021-68-9 CAPLUS
CN Pyrrolo[1,2-b]pyridazine-3-pentanoic acid, 4-(5-bromo-3-pyridinyl)-7-ethyl-2-[(1-pyrrolidinylcarbonyl)oxy]methyl-, ethyl ester (9CI) (CA INDEX NAME)



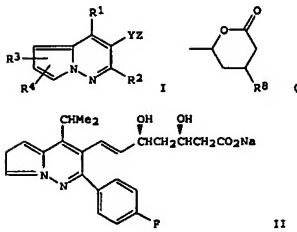
L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)



14 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2006 ACS ON STN
ACCESSION NUMBER: 1992:151782 CAPLUS
DOCUMENT NUMBER: 116-151782
TITLE: Preparation of pyrrolopyridazines as
hydroxymethylglutaryl (HMG) CoA reductase inhibitors
INVENTOR(S): Matsuo, Masaki; Manabe, Takashi; Okumura, Hirofumi;
Matsumura, Hiroshi; Fujii, Naoko
PATENT ASSIGNEE(S): Fujisawa Pharmaceutical Co., Ltd., Japan
SOURCE: PCT Int. Appl., 162 pp.
COPEN: PIXKD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9118903	A1	19911212	WO 1991-JP678	19910522
W: AU, CA, FI, HU, JP, KR, NO, SU, US				
RM: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, NL, SE				
AU 9178918	A1	19911231	AU 1991-78918	19910522
JP 06501918	T2	19940303	JP 1991-502910	19910522
CN 1056690	A	19911204	CN 1991-102784	19910524
PRIORITY APPLN. INFO.:			GB 1990-11337	A 19900522
			GB 1990-19173	A 19900903
			WO 1991-JP678	A 19910522

OTHER SOURCE(S): MARPAT 116:151782
GI



AB Title compds. I [R1, R2 = (C3-8 cycloalkyl) C1-6 alkyl, C3-8 cycloalkyl, (substituted) aryl; R3, R4 = H, (aryl)C1-6 alkyl, halo, (substituted)

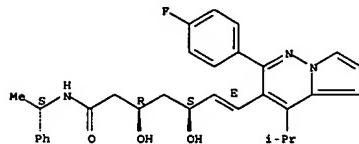
L4 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2006 ACS ON STN (Continued)
 aryl(thio), (protected) carboxy, acyl, (substituted) N-heterocycl.,
 etc.; Y = CH:CH, CH₂CH₂; Z = CHOCH₂CHOHCH₂RS, Q; R₅ = (protected) carboxy, RS
 - (protected OH) were prep'd. as inhibitors of HMG CoA reductase. Thus,
 MeCOCH₂CO₂Me was condensed with (E)-3-(2-(4-fluorophenyl)-4-
 isopropylpyrrolo[1,2-b]pyridazin-3-yl)acraldehyde (prep'n. given) in the
 presence of NaI to give (E)-RCH:CHCHOHCH₂CO₂Me [R =
 2-(4-fluorophenyl)-4-isopropylpyrrolo[1,2-b]pyridazin-3-yl]. This was
 reduced by Et₂Be²⁺ and NaBH₄ to give (±)-erythro-(E)-
 RCH:CHCHOHCH₂CHOHCH₂CO₂Me. Sapon. of the latter by NaOH gave title
 compd.
 II which had IC₅₀ of 0.009 µg/mL in vitro against HMG CoA reductase.

IT 139506-73-3P 139563-30-7P
PT: SPN (Synthetic preparation); PRRP (Preparation)

RE: SPN (Synthetic preparation); PREP (Preparation)
(preparation of, as hydroxymethylglutaryl CoA reductase inhibitor)

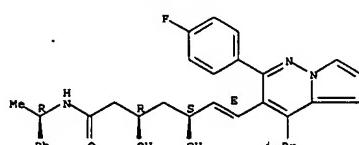
RN 139506-73-3 CAPLUS
CN 6-Heptenamide, 7-(2-(4-fluorophenyl)-4-(1-methylethyl)pyrrololo[1,2-
b]pyrazidin-3-yl)-3,5-dihydroxy-N-(1-phenylethyl)-, [3R-
1]-St1-3R-5S-, 6% fcr, (SCT). IUPAC INDEX NAME1

Absolute stereochemistry.
Double bond geometry as shown.



RN 139563-30-7 CAPLUS

CN 6-Heptenamide, 7-[2-(4-fluorophenyl)pyridazin-3-yl]-3,5-dihydroxy-



L4 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2006 ACS on STN (Continued)